

PCT

WORLD INTELLECTUAL PROPERTY ORGANIZATION  
International Bureau



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification <sup>6</sup> : <b>C11D 3/48, 3/22, 1/722</b>		A1	(11) International Publication Number: <b>WO 99/27060</b> (43) International Publication Date: <b>3 June 1999 (03.06.99)</b>
<p>(21) International Application Number: <b>PCT/US98/25148</b></p> <p>(22) International Filing Date: <b>24 November 1998 (24.11.98)</b></p> <p>(30) Priority Data: 08/979,730                    26 November 1997 (26.11.97)    US</p> <p>(71) Applicant (<i>for all designated States except US</i>): ALLERGAN SALES, INC. [US/US]; 2525 Dupont Drive, Irvine, CA 92612 (US).</p> <p>(72) Inventors; and (75) Inventors/Applicants (<i>for US only</i>): GRAHAM, Richard [US/US]; 5066 Balsawood, Irvine, CA 92612 (US). VEHIGE, Joseph, G. [US/US]; 28386 La Pradera, Laguna Niguel, CA 92677 (US).</p> <p>(74) Agents: FISHER, Carlos, A. et al.; Allergan Sales, Inc., 2525 Dupont Drive, Irvine, CA 92612 (US).</p>		<p>(81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).</p> <p><b>Published</b> <i>With international search report. Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.</i></p>	
<p>(54) Title: CONTACT LENS CLEANING COMPOSITIONS</p> <p>(57) Abstract</p> <p>Solutions useful to clean contact lenses include a surfactant component in an effective amount; and a viscosity inducing component, preferably selected from cellulosic derivatives and more preferably hydroxypropylmethyl cellulose, in an effective amount. Such solutions, which may include one or more additional components, have substantial contact lens cleaning benefits which, ultimately, lead to ocular health advantages and avoidance of problems caused by contact lens wear.</p>			

**FOR THE PURPOSES OF INFORMATION ONLY**

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	LS	Lesotho	SI	Slovenia
AM	Armenia	FI	Finland	LT	Lithuania	SK	Slovakia
AT	Austria	FR	France	LU	Luxembourg	SN	Senegal
AU	Australia	GA	Gabon	LV	Latvia	SZ	Swaziland
AZ	Azerbaijan	GB	United Kingdom	MC	Monaco	TD	Chad
BA	Bosnia and Herzegovina	GE	Georgia	MD	Republic of Moldova	TG	Togo
BB	Barbados	GH	Ghana	MG	Madagascar	TJ	Tajikistan
BE	Belgium	GN	Guinea	MK	The former Yugoslav Republic of Macedonia	TM	Turkmenistan
BF	Burkina Faso	GR	Greece			TR	Turkey
BG	Bulgaria	HU	Hungary	ML	Mali	TT	Trinidad and Tobago
BJ	Benin	IE	Ireland	MN	Mongolia	UA	Ukraine
BR	Brazil	IL	Israel	MR	Mauritania	UG	Uganda
BY	Belarus	IS	Iceland	MW	Malawi	US	United States of America
CA	Canada	IT	Italy	MX	Mexico	UZ	Uzbekistan
CF	Central African Republic	JP	Japan	NE	Niger	VN	Viet Nam
CG	Congo	KE	Kenya	NL	Netherlands	YU	Yugoslavia
CH	Switzerland	KG	Kyrgyzstan	NO	Norway	ZW	Zimbabwe
CI	Côte d'Ivoire	KP	Democratic People's Republic of Korea	NZ	New Zealand		
CM	Cameroon			PL	Poland		
CN	China	KR	Republic of Korea	PT	Portugal		
CU	Cuba	KZ	Kazakhstan	RO	Romania		
CZ	Czech Republic	LC	Saint Lucia	RU	Russian Federation		
DE	Germany	LI	Liechtenstein	SD	Sudan		
DK	Denmark	LK	Sri Lanka	SE	Sweden		
EE	Estonia	LR	Liberia	SG	Singapore		

CONTACT LENS CLEANING COMPOSITIONS

5

Background of the Invention

The present invention relates to compositions for treating, for example, cleaning, disinfecting, soaking, conditioning and wetting contact lenses. More 10 particularly, the invention relates to multi-purpose solutions useful in treating contact lenses, for example, for removing deposit material from contact lenses, for disinfecting contact lenses, for soaking, conditioning and/or wetting contact lenses and the like, which provide 15 substantial contact lens treating, e.g., cleaning, benefits to the users of such solutions.

Contact lenses need to be periodically treated, for example, cleaned, disinfected, soaked and the like, on a regular basis because of the tendency for a variety of 20 microbes and other materials to accumulate on the lenses and/or the need to provide the lenses in suitable condition for safe and comfortable wear.

Fu U.S. Patent 4,323,467 discloses aqueous compositions combining poly(oxyethylene)-poly(oxypropylene) substituted ethylenediamine surfactants, certain cellulose-derived polymer viscosity builders, germicidal agents, tonicity agents, sequestering agents and water for treating rigid contact lenses. The Fu patent does not disclose the use of hydroxypropylmethyl cellulose (HPMC) 30 or of any specific buffer.

British Patent 1,432,345 discloses a contact lens disinfecting composition including an ophthalmically acceptable biguanide in a total amount of from 0.0005% to 0.05% by weight. This British patent discloses that the 35 solution preferably has a pH of from 5 to 8 and employs a phosphate buffer. The patent also discloses employing additional bactericides, certain cellulose-derived thickening agents and non-ionic surfactants, as well as

disodium EDTA in concentrations of at least 0.1%. This patent does not disclose the use of HPMC.

Ogunbiyi et al U.S. Patent 4,758,595 discloses an aqueous solution of a biguanide in an amount of 0.000001 5 to 0.0003 weight percent in combination with a borate buffer system, EDTA, and one or more surfactants. This U.S. Patent additionally discloses that certain cellulose-derived viscosity builders can be included.

Mowrey-McKee et al U.S. Patent 5,422,073 discloses a 10 contact lens care solution including tromethamine, chelating agents, PHMB, surfactants and certain cellulose-derived viscosity inducing agents. This patent does not specifically disclose the use of HPMC.

There continues to be a need to provide new contact 15 lens treatment systems, for example, multi-purpose solutions, that provide one or more benefits, for example, more effective contact lens cleaning.

#### Summary of the Invention

20 New compositions for treating contact lenses have been discovered. The present compositions, for example, contact lens cleaning aqueous solutions and multi-purpose aqueous solutions, include surfactant components in amounts effective in removing deposit material from a 25 contact lens contacted with the composition, and effective amounts of viscosity inducing components, preferably HPMC in an amount in a range of about 0.05% to about 0.5% (w/v). The present HPMC-containing compositions preferably have increased or enhanced effectiveness in 30 removing deposit material from contact lenses contacted with the compositions relative to similar compositions without the HPMC. These compositions are surprising and unexpected in view of the above-noted prior art which discloses the use of cellulose-derived viscosity building 35 polymers other than HPMC. In addition, the present compositions preferably include antimicrobial components,

in combination with buffers to provide desired antimicrobial activity and performance effectiveness.

The inclusion of one or more still other components in the present compositions is effective in providing 5 additional beneficial properties to the compositions. The present compositions, in addition to being effective in cleaning contact lenses, preferably have a multitude of applications, for example, as disinfecting, soaking, wetting and conditioning compositions, for contact lens 10 care. The present compositions promote regular and consistent contact lens care and, ultimately, lead to or facilitate better ocular health.

Any suitable, preferably ophthalmically acceptable, surfactant component which is effective in cleaning 15 contact lenses may be employed. The surfactant component preferably is nonionic and, more preferably, is selected from 4-(1,1,3,3-tetramethylbutyl)phenol/poly(oxyethylene) polymers, poly(oxyethylene)-poly(oxypropylene) block copolymers and mixtures thereof.

20 Although any suitable, for example, ophthalmically acceptable, viscosity inducing or thickening agent may be included in the present compositions, the viscosity inducing component preferably is selected from cellulosic derivatives and mixtures thereof, and more preferably is 25 HPMC. The viscosity inducing component preferably is present in an amount in the range of about 0.05% to about 0.5% (w/v). Without wishing to limit the invention to any particular theory of operation, it is believed that the presence of a viscosity inducing component at least 30 assists in providing the present compositions with enhanced passive contact lens cleaning properties. Passive cleaning refers to the cleaning which occurs during soaking of a contact lens, without mechanical or enzymatic enhancement. In particular, it has unexpectedly 35 been found that the present compositions with HPMC present are more effective in passive cleaning of contact lenses

relative to similar compositions without HPMC. The present combinations of components, including such viscosity inducing components, are effective in providing the degree of enhanced contact lens cleaning described  
5 herein.

In one embodiment of the present invention, multi-purpose solutions for contact lens care are provided. Such solutions comprise an aqueous liquid medium; a non-oxidative antimicrobial component in an amount effective  
10 to disinfect a contact lens contacted with the solution; a surfactant in an amount effective in cleaning a contact lens contacted with the solution; a buffer component, preferably a phosphate buffer component in an amount effective in maintaining the pH of the solution within a  
15 physiologically acceptable range; a viscosity inducing component preferably HPMC, present in an effective amount; and a tonicity component in an amount effective in providing the desired tonicity to the solution.

The antimicrobial component may be any suitable, preferably ophthalmically acceptable, material effective to disinfect a contact lens contacted with the present solutions. In one embodiment, the antimicrobial component is non-oxidative. Preferably, the non-oxidative antimicrobial component is selected from biguanides,  
25 biguanides polymers, salts thereof and mixtures thereof, and is present in an amount in the range of about 0.1 ppm to about 3 ppm or less than 5 ppm (w/v). The preferred relatively reduced concentration of the antimicrobial component has been found to be very effective, in the  
30 present compositions, in disinfecting contact lenses contacted with the compositions, while at the same time promoting lens wearer/user comfort and acceptability.

Although any suitable, preferably ophthalmically acceptable, tonicity component may be employed, a very  
35 useful tonicity component is sodium chloride or a combination of sodium chloride and potassium chloride.

The present compositions preferably include an effective amount of a chelating component. Any suitable, preferably ophthalmically acceptable, chelating component may be included in the present compositions, although 5 ethylenediaminetetraacetic acid (EDTA), salts thereof and mixtures thereof are particularly effective.

Various combinations of two or more of the above-noted components may be used in providing at least one of the benefits described herein. Therefore, each and every 10 such combination is included within the scope of the present invention.

These and other aspects of the present invention are apparent in the following detailed description, examples and claims.

15

#### Detailed Description of the Invention

The present invention is directed to solutions useful for cleaning contact lenses and to multi-purpose solutions useful for treating, for example, cleaning, disinfecting, 20 soaking, rinsing, wetting, conditioning and the like, contact lenses. Any contact lenses, for example, conventional hard contact lenses, rigid gas permeable contact lenses and soft, hydrophilic or hydrogel, contact lenses, can be treated in accordance with the present 25 invention.

The present compositions, preferably solutions, useful for cleaning a contact lens comprise an aqueous liquid medium, a surfactant component in an amount effective in removing deposit material from a contact lens contacted with the composition, and an effective amount of 30 a viscosity inducing component, preferably HPMC in an amount in the range of about 0.05% to about 0.5% (w/v).

In one embodiment, the present compositions, preferably solutions, comprise a liquid aqueous medium; a 35 non-oxidative antimicrobial component in the liquid aqueous medium in an amount effective to disinfect a

contact lens contacted with the composition; a surfactant, preferably a nonionic surfactant, component in an amount effective in cleaning, or removing deposit material from, a contact lens contacted with the composition; a buffer 5 component, for example, a phosphate buffer component, in an amount effective in maintaining the pH of the composition within a physiologically acceptable range; an effective amount of a viscosity inducing component, preferably HPMC; and an effective amount of a tonicity 10 component.

The present compositions preferably include an effective amount of a chelating or sequestering component, more preferably in a range of less than 0.1% (w/v). Each 15 of the components, in the concentration employed, included in the compositions and the formulated compositions of the present invention preferably are ophthalmically acceptable. In addition, each of the components, in the concentration employed, included in the present compositions preferably is soluble in the liquid aqueous 20 medium.

A composition or component thereof is "ophthalmically acceptable" when it is compatible with ocular tissue, that is, it does not cause significant or undue detrimental effects when brought into contact with ocular tissue. 25 Preferably, each component of the present compositions is also compatible with the other components of the present compositions.

The surfactant component is present in an amount effective in cleaning, that is to at least facilitate 30 removing, and preferably effective to remove, debris or deposit material from, a contact lens contacted with the surfactant-containing solution. Exemplary surfactant components include, but are not limited to, nonionic surfactants, for example, polysorbates (such as 35 polysorbate 20-Trademark Tween 20), 4-(1, 1, 3, 3-tetramethylbutyl) phenol polymers (such as the polymer

sold under the trademark Tyloxapol), poly(oxyethylene)-poly(oxypropylene) block copolymers, glycolic esters of fatty acids and the like, and mixtures thereof.

The surfactant component more preferably is nonionic,  
5 and still more preferably is selected from 4-(1,1,3,3-tetrabutyl)phenol/poly(oxyethylene) polymers, poly(oxyethylene)-poly(oxypropylene) block copolymers and mixtures thereof. Such block copolymers can be obtained commercially from the BASF Corporation under the trademark  
10 Pluronic<sup>7</sup>, and can be generally described as polyoxyethylene/polyoxypropylene condensation polymers terminated in primary hydroxyl groups. They may be synthesized by first creating a hydrophobe of desired molecular weight by the controlled addition of propylene  
15 oxide to the two hydroxyl groups of propylene glycol. In the second step of the synthesis, ethylene oxide is added to sandwich this hydrophobe between hydrophile groups.

In accordance with a more preferred embodiment of the invention, such block copolymers having molecular weights  
20 in the range of about 2500 to 13,000 daltons are suitable, with a molecular weight range of about 6000 to about 12,000 daltons being still more preferred. Specific examples of surfactants which are satisfactory include:  
25 poloxamer 108, poloxamer 188, poloxamer 237, poloxamer 238, poloxamer 288, poloxamer 407.

The amount of surfactant component present varies over a wide range depending on a number of factors, for example, the specific surfactant or surfactants being used, the other components in the composition and the  
30 like. Often the amount of surfactant is in the range of about 0.005% or about 0.01% to about 0.1% or about 0.5% or about 0.8% (w/v).

The viscosity inducing component is effective to enhance and/or prolong the cleaning and wetting activity  
35 of the surfactant component and/or condition the lens surface rendering it more hydrophilic (less lipophilic)

and/or to act as a demulcent on the eye. Increasing the solution viscosity provides a film on the lens which may facilitate comfortable wearing of the treated contact lens. The viscosity inducing component may also act to 5 cushion the impact on the eye surface during insertion and serves also to alleviate eye irritation.

Suitable viscosity inducing components include, but are not limited to, water soluble natural gums, cellulose-derived polymers and the like. Useful natural gums 10 include guar gum, gum tragacanth and the like. Useful cellulose-derived viscosity inducing components include cellulose-derived polymers, such as hydroxypropyl cellulose, HPMC, carboxymethyl cellulose, methyl cellulose, hydroxyethyl cellulose and the like. More 15 preferably, the viscosity inducing agent is selected from cellulose derivatives (polymers) and mixtures thereof.

A very useful viscosity inducing component is HPMC. The viscosity inducing component, and in particular HPMC, has been found to enhance the ability of the present 20 compositions in cleaning, for example, in passively cleaning (e.g., without manual rubbing), contact lenses.

The viscosity inducing component is used in an amount effective to increase the viscosity of the solution, preferably to a viscosity in the range of about 1.5 to 25 about 30, or even as high as about 750, cps at 25°C, preferably as determined by USP test method No. 911 (USP 23, 1995). The amount of viscosity inducing component preferably is in the range of about 0.01% to about 5% (w/v), with amounts of about 0.05% to about 0.5% being 30 more preferred.

The present compositions preferably further comprise effective amounts of one or more additional components, such as an antimicrobial component; a buffer component; a chelating or sequestering component; a tonicity component; 35 and the like and mixtures thereof. The additional component or components may be selected from materials

- which are known to be useful in contact lens care compositions and are included in amounts effective to provide the desired effect or benefit. When an additional component is included, it is preferably compatible under typical use and storage conditions with the other components of the composition. For instance, the aforesaid additional component or components preferably are substantially stable in the presence of the surfactant and viscosity inducing components described herein.
- The presently useful antimicrobial components include chemicals which derive their antimicrobial activity through a chemical or physiochemical interaction with microbes or microorganisms, such as those contaminating a contact lens. Suitable antimicrobial components are those generally employed in ophthalmic applications and include, but are not limited to, quaternary ammonium salts used in ophthalmic applications such as poly[dimethylimino-2-butene-1,4-diyl] chloride, alpha-[4-tris(2-hydroxyethyl)ammonium]-dichloride (chemical registry number 75345-27-6, available under the trademark Polyquaternium 17 from Onyx Corporation), tromethamine, benzalkonium halides, and biguanides, such as salts of alexidine, alexidine-free base, salts of chlorhexidine, hexamethylene biguanides and their polymers, and salts thereof, antimicrobial polypeptides, chlorine dioxide precursors, and the like and mixtures thereof. Generally, the hexamethylene biguanide polymers (PHMB), also referred to as polyaminopropyl biguanide (PAPB), have molecular weights of up to about 100,000. Such biguanide polymers are known and are disclosed in Ogunbiyi et al U.S. Patent No. 4,758,595, the disclosure of which is hereby incorporated in its entirety by reference herein.
- The antimicrobial components useful in the present invention preferably are present in the liquid aqueous medium in concentrations in the range of about 0.00001% to about 2% (w/v).

More preferably, the antimicrobial component is present in the liquid aqueous medium at an ophthalmically acceptable or safe concentration such that the user can remove the disinfected lens from the liquid aqueous medium 5 and thereafter directly place the lens in the eye of safe and comfortable wear.

The antimicrobial components suitable for inclusion in the present invention include chlorine dioxide precursors. Specific examples of chlorine dioxide 10 precursors include stabilized chlorine dioxide (SCD), metal chlorites, such as alkali metal and alkaline earth metal chlorites, and the like and mixtures thereof. Technical grade sodium chlorite is a very useful chlorine dioxide precursor. Chlorine dioxide-containing complexes, 15 such as complexes of chlorine dioxide with carbonate, chlorine dioxide with bicarbonate and mixtures thereof are also included as chlorine dioxide precursors. The exact chemical composition of many chlorine dioxide precursors, for example, SCD and the chlorine dioxide complexes, is 20 not completely understood. The manufacture or production of certain chlorine dioxide precursors is described in McNicholas U.S. Patent 3,278,447, which is incorporated in its entirety herein by reference. Specific examples of useful SCD products include that sold under the trademark 25 Dura Klor by Rio Linda Chemical Company, Inc., and that sold under the trademark Anthium Dioxide by International Dioxide, Inc.

If a chlorine dioxide precursor is included in the present compositions, it preferably is present in an 30 effective contact lens disinfecting amount. Such effective disinfecting concentrations preferably are in the range of about 0.002 to about 0.06% (w/v) of the present compositions. Such chlorine dioxide precursors may be used in combination with other antimicrobial 35 components, such as biguanides, biguanide polymers, salts thereof and mixtures thereof.

In the event that chlorine dioxide precursors are employed as antimicrobial components, the compositions preferably have an osmolality of at least about 200 mOsmol/kg and are buffered to maintain the pH within an acceptable physiological range, for example, a range of about 6 to about 10.

It has been found that reduced amounts of non-oxidative antimicrobial components, for example, in a range of about 0.1 ppm to about 3 ppm or less than 5 ppm (w/v), in the present compositions are effective in disinfecting contact lenses and reduce the risk of such antimicrobial components causing ocular discomfort and/or irritation. Such reduced concentration of antimicrobial component is very useful when the antimicrobial component employed is selected from biguanides, biguanide polymers, salts thereof and mixtures thereof.

When a contact lens is desired to be disinfected by the present compositions, an amount of the antimicrobial component effective to disinfect the lens is used. Preferably, such an effective amount of the antimicrobial component reduces the microbial burden or load on the contact lens by one log order in three hours. More preferably, an effective amount of the disinfectant reduces the microbial load by one log order in one hour.

The buffer component is present in an amount effective to maintain the pH of the composition or solution in the desired range, for example, in a physiologically acceptable range of about 4 or about 5 or about 6 to about 8 or about 9 or about 10. In particular, the solution preferably has a pH in the range of about 6 to about 8. Any material which is ophthalmically acceptable and has buffering effectiveness in the present applications may be employed. Such buffers may include organic materials, such as tromethamine and the like, inorganic materials, such as phosphates, borates carbonates and the like, and mixtures thereof.

Particularly useful phosphate buffer components include one or more phosphate buffers, for example, combinations of monobasic phosphates, dibasic phosphates and the like, such as those selected from phosphate salts of alkali 5 and/or alkaline earth metals. Examples of suitable phosphate buffers include one or more of sodium dibasic phosphate ( $\text{Na}_2\text{HPO}_4$ ), sodium monobasic phosphate ( $\text{NaH}_2\text{PO}_4$ ) and potassium monobasic phosphate ( $\text{KH}_2\text{PO}_4$ ). The present buffer components frequently are used in amounts in a 10 range of about 0.01% or about 0.02% to about 1% or about 2% (w/v) or more.

A chelating or sequestering component preferably is included in an amount effective to enhance the effectiveness of the antimicrobial component and/or to 15 complex with metal ions to provide more effective cleaning of the contact lens.

A wide range of organic acids, amines or compounds which include an acid group and an amine function are capable of acting as chelating components in the present 20 compositions. For example, nitrilotriacetic acid, diethylenetriaminepentacetic acid, hydroxyethylethylene-diaminetriacetic acid, 1,2-diaminocyclohexane tetraacetic acid, hydroxyethylaminodiacetic acid, ethylenediaminetetraacetic acid and its salts, polyphosphates, citric 25 acid and its salts, tartaric acid and its salts, and the like and mixtures thereof, are useful as chelating components. Ethylenediaminetetraacetic acid (EDTA) and its alkali metal salts, are preferred, with disodium salt of EDTA, also known as disodium edetate, being 30 particularly preferred.

The chelating component preferably is present in an effective amount, for example, in a range of about 0.01% and about 1% (w/v) of the solution.

In a very useful embodiment, particularly when the 35 chelating component is EDTA, salts thereof and mixtures thereof, a reduced amount is employed, for example, in the

range of less than about 0.1% (w/v). Such reduced amounts of chelating component have been found to be effective in the present compositions while, at the same time, providing for reduced discomfort and/or ocular irritation.

5       The liquid aqueous medium used is selected to have no substantial deleterious effect on the lens being treated, or on the wearer of the treated lens. The liquid medium is constituted to permit, and even facilitate, the lens treatment or treatments by the present compositions. The  
10      liquid aqueous medium advantageously has an osmolality in the range of at least about 200 mOsmol/kg for example, about 300 or about 350 to about 400 mOsmol/kg. The liquid aqueous medium more preferably is substantially isotonic or hypertonic (for example, slightly hypertonic) and/or is  
15      ophthalmically acceptable.

The liquid aqueous medium preferably includes an effective amount of a tonicity component to provide the liquid medium with the desired tonicity. Such tonicity components may be present in the liquid aqueous medium  
20      and/or may be introduced into the liquid aqueous medium. Among the suitable tonicity adjusting components that may be employed are those conventionally used in contact lens care products, such as various inorganic salts. Sodium chloride and/or potassium chloride and the like are very  
25      useful tonicity components. The amount of tonicity component included is effective to provide the desired degree of tonicity to the solution. Such amount may, for example, be in the range of about 0.4% to about 1.5% (w/v). If a combination of sodium chloride and potassium  
30      chloride is employed, it is preferred that the weight ratio of sodium chloride to potassium chloride be in the range of about 3 to about 6 or about 8.

Methods for treating a contact lens using the herein described compositions are included within the scope of  
35      the invention. Such methods comprise contacting a contact

lens with such a composition at conditions effective to provide the desired treatment to the contact lens.

The contacting temperature is preferred to be in the range of about 0°C to about 100°C, and more preferably in 5 the range of about 10°C to about 60°C and still more preferably in the range of about 15°C to about 30°C. Contacting at or about ambient temperature is very convenient and useful. The contacting preferably occurs at or about atmospheric pressure. The contacting 10 preferably occurs for a time in the range of about 5 minutes or about 1 hour to about 12 hours or more.

The contact lens can be contacted with the liquid aqueous medium by immersing the lens in the medium. During at least a portion of the contacting, the liquid 15 medium containing the contact lens can be agitated, for example, by shaking the container containing the liquid aqueous medium and contact lens, to at least facilitate removal of deposit material from the lens. After such contacting step, the contact lens may be manually rubbed 20 to remove further deposit material from the lens. The cleaning method can also include rinsing the lens substantially free of the liquid aqueous medium prior to returning the lens to a wearer's eye.

The following non-limiting examples illustrate 25 certain aspects of the present invention.

#### EXAMPLE 1

A solution is prepared by blending together the 30 following components:

	PHMB (polyhexamethylene biguanide)	1 ppm (w/v)
35	Disodium EDTA	0.05% (w/v)
	Tyloxapol	0.025% (w/v)
	Tromethamine	1.2% (w/v)

	HPMC (Hydroxypropylmethyl Cellulose)	0.15% (w/v)
5	Sodium Chloride	0.37% (w/v)
	Water (USP)	Q.S. 100%
10	pH (adjusted with HCl)	7.5

Approximately three (3)ml of this solution is introduced into a lens vial containing a lipid, oily deposit laden, hydrophilic or soft contact lens. The contact lens is maintained in this solution at room temperature for at least about four (4) hours. This treatment is effective to disinfect the contact lens. In addition, it is found that a substantial portion of the deposits previously present on the lens has been removed. This demonstrates that this solution has substantial passive contact lens cleaning ability.

After this time, the lens is removed from the solution and is placed in the lens wearer's eye for safe and comfortable wear. Alternately, after the lens is removed from the solution, it is rinsed with another quantity of this solution and the rinsed lens is then placed in the lens wearer's eye for safe and comfortable wear.

#### EXAMPLE 2

Example 1 is repeated except that the lens is rubbed and rinsed with a different quantity of the solution prior to being placed in the lens vial. After at least about four (4) hours, the lens is removed from the solution. The lens is then placed in the lens wearer's eye for safe and comfortable wear.

EXAMPLE 3

The solution of Example 1 is used as a long-term soaking medium for a hydrophilic contact lens. Thus,  
5 approximately three (3) ml of this solution is placed in a vial and a contact lens is maintained in the solution at room temperature for about sixty (60) hours. After this soaking period, the lens is removed from the solution and placed in the lens wearer's eye for safe and comfortable  
10 wear. Alternately, after the lens is removed from the solution, it is rinsed with another quantity of this solution and the rinsed lens is then placed in the lens wearer's eye for safe and comfortable wear.

15

EXAMPLE 4

A hydrophilic contact lens is ready for wear. In order to facilitate such wearing, one or two drops of the solution of Example 1 is placed on the lens immediately  
20 prior to placing the lens in the lens wearer's eye. The wearing of this lens is comfortable and safe.

EXAMPLE 5

25 A lens wearer wearing a contact lens applies one or two drops of the solution of Example 1 in the eye wearing the lens. This effects a re-wetting of the lens and provides for comfortable and safe lens wear.

30

EXAMPLE 6

A series of tests are conducted to evaluate the passive contact lens cleaning ability of the solution prepared in accordance with Example 1 compared to other  
35 solutions.

The first of these other solutions, referred to hereinafter as Composition A, is similar to the solution prepared in accordance with Example 1 except no HPMC is included.

5       The second of these other solutions, referred to hereinafter as Composition B, is sold under the trademark ReNu7 by Bausch & Lomb and includes 0.5 ppm PHMB, a poly(oxyethylene)-poly(oxypropylene) substituted ethylenediamine surfactant, a borate buffer system, 0.1%  
10 disodium EDTA, and sodium chloride as a tonicity agent.

The remaining other solutions are as follows:

15      Composition C is sold by Alcon under the trademark Opti-Free™

Composition D is sold by Ciba Vision Care under the trademark Solo Care™soft

Composition E a saline solution sold by Allergan under the trademark Lens Plus™

20      Each of these compositions is tested to evaluate its passive cleaning ability, specifically its ability to passively remove lipid-containing soil from a contact lens.

25      These tests are conducted as follows. A model lipid soil is prepared by combining one part by weight of Apiezon AP 101, 1.38 parts by weight paraffin oil and 0.01 parts by weight of Oil Red O. A red grease mixture is produced. This soil is deposited by first coating a  
30 circular stamp device with a diameter of about 1/2 inch which is plugged with cotton. The coated device is then stamped on the bottom of a tissue culture well made of polystyrene making sure that a light uniform coat is deposited on the bottom surface. Three (3) wells are  
35 coated for each solution to be tested. Two (2) sets of the coated wells are prepared. One set for one hour soaking and the second set for four (4) hours soaking. The coated wells are photographed in a photocopy machine and marked as the initial point.

The plates are cleaned as follows. 10ml of each of the cleaning solutions is pipetted into the freshly prepared coated wells. One set of wells is allowed to soak for one hour and the second set is allowed to soak 5 for four (4) hours. After the soaking cycle, the solution is decanted by flipping the well upside down.

Visual observations of any changes in the coating are made during the soaking cycle. At the end of the soaking cycle the wells are again photographed in a photocopy 10 machine.

The estimated passive cleaning resulting from the soaking is ranked 1 to 5 with 1 representing the highest degree of passive cleaning and 5 representing the lowest degree of passive cleaning. The results of this ranking 15 are as follows.

	<u>Solution</u>	<u>1 hr. Soaking</u>	<u>4 hrs. Soaking</u>	<u>Average Rank</u>
20	Example 1	1	1	1
	Composition A	2	2	2
	Composition B	4	4	4
	Composition C	5	5	5
	Composition D	3	3	3
25	Composition E	5	5	5

After the one (1) hour soak, dispersion and solubilization of some of the lipid substance is observed in the wells soaked with the solution in accordance with 30 Example 1, Composition A and Composition D. Strings of the coating start to roll up and are seen floating on the surface of the solution. These results are seen again after the four (4) hour soak.

These results indicate that the solution in 35 accordance with Example 1 is the most effective in passive cleaning regimens both in the one (1) hour and four (4) hours soaking. Visual observations show the effectiveness rankings of the Example 1 solution and Compositions A and B to be: Example 1 > Composition A >> Composition B. 40 Composition C is the least efficacious of the solutions,

its lipid cleaning efficacy comparable only to the saline solution, Composition E. Composition A, on the other hand, shows more cleaning after the four (4) hours soaking period, while after one (1) hour soaking showing only the 5 beginnings of dispersion of the coating. Composition D is a more effective passive cleaner than is Composition B.

When comparing the solution in accordance with Example 1 with Composition A, it is seen that the inclusion of HPMC in the solution of Example 1, in 10 combination with the other ingredients present, provides an enhancement in passive cleaning efficacy.

While this invention has been described with respect to various specific examples and embodiments, it is to be understood that the invention is not limited thereto and 15 that it can be variously practiced within the scope of the following claims.

WHAT IS CLAIMED IS:

1. A composition for cleaning a contact lens comprising:

an aqueous liquid medium;

a surfactant component in an amount effective in removing deposit material from a contact lens contacted with said composition; and

hydroxypropylmethyl cellulose in an amount in a range of about 0.05% to about 0.5% (w/v).

2. The composition of claim 1 which has enhanced effectiveness in removing deposit material from a contact lens contacted with said composition relative to a similar composition without the hydroxypropylmethyl cellulose.

3. The composition of claim 1 which has enhanced effectiveness in passively removing deposit material from a contact lens contacted with said composition relative to a similar composition without the hydroxypropylmethyl cellulose.

4. The composition of claim 1 wherein said surfactant is present in an amount in the range of about 0.01% to about 0.8% (w/v).

5. The composition of claim 1 wherein said surfactant component is selected from the group consisting of nonionic surfactants and mixtures thereof.

6. The composition of claim 1 wherein said surfactant component is selected from the group consisting of polysorbates, 4-(1,1,3,3-tetramethylbutyl) phenol/poly(oxyethylene) polymers, poly(oxyethylene)-poly(oxypropylene) block copolymers, glycolic esters of fatty acids, alkyl ether sulfates and mixtures thereof.

7. The composition of claim 1 wherein said surfactant component is selected from the group consisting of 4-(1,1,3,3-tetramethylbutyl) phenol/poly(oxyethylene) polymers, poly(oxyethylene)-poly(oxypropylene) block copolymers and mixtures thereof.

8. The composition of claim 1 wherein said surfactant component is selected from the group consisting of 4-(1,1,3,3-tetrabutyl) phenol/poly(oxyethylene) polymers and mixtures thereof.

9. The composition of claim 1 which includes a buffer component in an amount effective in maintaining the pH of said composition within a physiologically acceptable range, and a tonicity component in an amount effective in providing the desired tonicity to said composition.

10. The composition of claim 1 which further comprises an effective amount of a chelating component.

11. The composition of claim 1 which further comprises an antimicrobial component in an amount effective to disinfect a contact lens contacted with said composition.

12. The composition of claim 11 wherein said antimicrobial component is a non-oxidative antimicrobial component.

13. The composition of claim 11 wherein said antimicrobial component is selected from the group consisting of biguanides, biguanide polymers, salts thereof and mixtures thereof.

14. A composition for cleaning a contact lens comprising:

an aqueous liquid medium;

a non-oxidative antimicrobial component in an effective amount of less than 5 ppm;

a surfactant component selected from the group consisting of 4-(1,1,3,3-tetramethylbutyl) phenol/poly(oxyethylene) polymers, poly(oxyethylene)-poly(oxyethylene) block copolymers and mixtures thereof in an amount effective in removing deposit material from a contact lens contacted with said composition;

a phosphate buffer component in an amount effective in maintaining the pH of said solution within a physiologically acceptable range; and

a viscosity inducing component in an amount effective in increasing the viscosity of said composition, provided that said composition has increased effectiveness in removing deposit material from a contact lens contacted with said composition relative to a similar composition without said viscosity inducing component.

15. The composition of claim 14 which has enhanced effectiveness in passively removing deposit material from a contact lens contacted with said composition relative to a similar composition without the viscosity inducing component.

16. The composition of claim 14 wherein said viscosity inducing component selected from the group consisting of cellulosic derivatives and mixtures thereof and being present in an effective amount in a range of about 0.05% to about 0.5% (w/v).

17. The composition of claim 14 wherein said viscosity inducing component is hydroxypropylmethyl cellulose.

18. The composition of claim 14 wherein said non-oxidative antimicrobial component is selected from the group consisting of biguanides, biguanide polymers, salts thereof and mixtures thereof.

19. The composition of claim 14 wherein said non-oxidative antimicrobial component is selected from the group consisting of polyhexamethylene biguanide, salts thereof and mixtures thereof.

20. The composition of claim 14 wherein said surfactant component is present in an amount in a range of about 0.01% to about 0.8% (w/v).

# INTERNATIONAL SEARCH REPORT

International Application No.  
PCT/US 98/25148

## A. CLASSIFICATION OF SUBJECT MATTER

C 11 D 3/48, C 11 D 3/22, C 11 D 1/722

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

C 11 D

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 3311577 A (RANKIN, B.F.) 28 March 1967 (28.03.67), the whole document. --	1-11, 14-17
X	Derwent Accession, week 8832, London: Derwent Publications Ltd., AN 88-224463 (25), Class C11D; & JP,A,63-159821 (TOME SANGYO KK), abstract. --	1-11, 14-17
A, P	WO 98/32421 A1 (BAUSCH & LOMB INCORPORATED) 30 July 1998 (30.07.98), examples, claims 1-3. --	1-11, 14-17
A	US 5591426 A (DABROWSKI et al.) 07 January	1-3, 14-17

Further documents are listed in the continuation of box C.

Patent family members are listed in annex.

### \* Special categories of cited documents :

- "A" document defining the general state of the art which is not considered to be of particular relevance
- "E" earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- "O" document referring to an oral disclosure, use, exhibition or other means
- "P" document published prior to the international filing date but later than the priority date claimed

- "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- "&" document member of the same patent family

Date of the actual completion of the international search

15 March 1999

Date of mailing of the international search report

06.05.1999

Name and mailing address of the ISA

European Patent Office, P.O. 5818 Patentam 2  
NL - 2280 HV Rijswijk  
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  
Fax (+31-70) 340-3016

Authorized officer

SEIRFAI e.h.

# INTERNATIONAL SEARCH REPORT

International Application No.  
PCT/US 98/25148

## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
	<p>1997 (07.01.97), the whole document. -----</p>	